## RECEIVED

## SEQUENCE LISTING

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<120> BIOSYNTHESIS GENES & TRANSFER OF 6-DESOXYHEXOSES IN SACCHAROPOLYSPORA ERYTHRAEA AND IN STREPTOMYCES ANTIBIOTICUS AND THEIR USE

<130> 146.1335

<140> 09/463705 <141> 2000-02-23

<150> PCT/FR98/01593

<151> 1998-07-21

<150> 98/07411

<151> 1988-06-12

<150> 97/09458

<151> 1997-07-25

<160> 61

<170> PatentIn Ver. 2.1

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Ile Asn Cys Leu Asp Thr Ala Asp Met Tyr Gly Trp Arg Leu Tyr Lys
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Cys Glu Gly Ser Leu Arg Arg Leu Gly Val Asp His Ile Asp Val Tyr 115 120 125

Gln Met His His Ile Asp Arg Ser Ala Pro Trp Asp Glu Val Trp Gln 130 135 140

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Ser Asn Phe Ala Gly Trp His Ile Ala Ala Ala Gln Glu Asn Ala Ala 165 170 175

Arg Arg His Ser Leu Gly Met Val Ser His Gln Cys Leu Tyr Asn Leu 180 185 190

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Gly Leu Gly Val Phe Ala Trp Ser Pro Leu His Gly Gly Leu Leu Ser 210 215 220

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Asp His Ala Thr Ala Arg Gln Val Leu Asp Asp Pro Ala Phe Thr Arg 65 70 75 80

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Ala Glu Trp Ala Gln Pro Phe Arg Asp Val His Ala Ala Ser Trp Glu 100 105 110

Gly Glu Val Pro Asp Val Gly Glu Leu Ala Glu Ser Phe Ala Gly Leu 115 120 125

Leu Pro Gly Ala Gly Ala Arg Leu Asp Leu Val Gly Asp Phe Ala Trp 130 135 140

Gln Val Pro Val Gln Gly Met Thr Ala Val Leu Gly Ala Ala Gly Val 145 150 155 160

Leu Arg Gly Ala Ala Trp Asp Ala Arg Val Ser Leu Asp Ala Gln Leu 165 170 175

Ser Pro Gln Gln Leu Ala Val Thr Glu Ala Ala Val Ala Leu Pro 180 185 190

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Thr Ala Glu Val Arg Leu Gly Glu His Val Ile Gly Glu Glu Glu Glu 260 265 . 270

Val Val Val Val Ala Ala Ala Asn Arg Asp Pro Glu Val Phe Ala 275 280 285

Glu Pro Asp Arg Leu Asp Val Asp Arg Pro Asp Ala Asp Arg Ala Leu 290 295 300

Ser Ala His Arg Gly His Pro Gly Arg Leu Glu Glu Leu Val Thr Ala 305 310 315 320

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Glu Arg Asp Pro Ala Thr Leu Thr Trp Glu His Leu Leu Gly Met Gln 85 90 95

Thr Val Leu Thr Pro Thr Phe Tyr Ala Leu Met Ser Pro Asp Thr Leu

Ile Glu Gly Met Val Ser Phe Cys Arg Lys Trp Arg Pro Asp Leu Val 120 Ile Trp Glu Pro Leu Thr Phe Ala Ala Pro Ile Ala Ala Val Thr 135 Gly Thr Pro His Ala Arg Leu Leu Trp Gly Pro Asp Ile Thr Thr Arg 150 Ala Arg Gln Asn Phe Leu Gly Leu Leu Pro Asp Gln Pro Glu Glu His 170 Arg Glu Asp Pro Leu Ala Glu Trp Leu Thr Trp Thr Leu Glu Lys Tyr Gly Gly Pro Ala Phe Asp Glu Glu Val Val Gly Gln Trp Thr Ile Asp Pro Ala Pro Ala Ala Ile Arg Leu Asp Thr Gly Leu Lys Thr Val Gly Met Arg Tyr Val Asp Tyr Asn Gly Pro Ser Val Val Pro Glu Trp 235 Leu His Asp Glu Pro Glu Arg Arg Val Cys Leu Thr Leu Gly Ile 250 Ser Ser Arg Glu Asn Ser Ile Gly Gln Val Ser Ile Glu Glu Leu Leu 265 Gly Ala Val Gly Asp Val Asp Ala Glu Ile Ile Ala Thr Phe Asp Ala 280 Gln Gln Leu Glu Gly Val Ala Asn Ile Pro Asp Asn Val Arg Thr Val 295 Gly Phe Val Pro Met His Ala Leu Leu Pro Thr Cys Ala Ala Thr Val 305 310 His His Gly Gly Pro Gly Ser Trp His Thr Ala Ala Ile His Gly Val 330 Pro Gln Val Ile Leu Pro Asp Gly Trp Asp Thr Gly Val Arg Ala Gln Arg Thr Gln Glu Phe Gly Ala Gly Ile Ala Leu Pro Val Pro Glu Leu Thr Pro Asp Gln Leu Arg Glu Ser Val Lys Arg Val Leu Asp Asp Pro

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Val Arg Asp Arg Leu Pro Ser Ala Ser Ser Leu Leu Asp Val Ala Cys 770  ggg acc ggc acc cac ctg cgc cgg ttc gcc gac ctc ttc gac gac gtg Gly Thr Gly Thr His Leu Arg Arg Phe Ala Asp Leu Phe Asp Asp Val 785  acc ggg ctg gag ctg tcg gcg gcg atg atc gag gtc gcc cgg ccg cag Thr Gly Leu Glu Leu Ser Ala Ala Met Ile Glu Val Ala Arg Pro Gln 805  ctc ggc ggc atc ccg gtg ctg cag ggc gac atg cgc gac ttc gcg ctg Leu Gly Gly Ile Pro Val Leu Gln Gly Asp Met Arg Asp Phe Ala Leu 820  gat cgc gag ttc gac gcc gtc acc tgc atg ttc agc tcc atc ggg cac Asp Arg Glu Phe Asp Ala Val Thr Cys Met Phe Ser Ser Ile Gly His 835	698 746 794

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aac ggc acg cgc o Asn Gly Thr Arg I 1160		Ile Leu Val			910
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Met Arg Val Leu Leu Thr Ser Phe Ala His Arg Thr His Phe Gln Gly

- 21 -

Leu Val Pro Leu Ala Trp Ala Leu Arg Thr Ala Gly His Asp Val Arg Val Ala Ala Gln Pro Ala Leu Thr Asp Ala Val Ile Gly Ala Gly Leu 40 Thr Ala Val Pro Val Gly Ser Asp His Arg Leu Phe Asp Ile Val Pro Glu Val Ala Ala Gln Val His Arg Tyr Ser Phe Tyr Leu Asp Phe Tyr 70 75 His Arg Glu Gln Glu Leu His Ser Trp Glu Phe Leu Leu Gly Met Gln Glu Ala Thr Ser Arg Trp Val Tyr Pro Val Val Asn Asn Asp Ser Phe Val Ala Glu Leu Val Asp Phe Ala Arg Asp Trp Arg Pro Asp Leu Val Leu Trp Glu Pro Phe Thr Phe Ala Gly Ala Val Ala Ala Arg Ala Cys 135 Gly Ala Ala His Ala Arg Leu Leu Trp Gly Ser Asp Leu Thr Gly Tyr Phe Arg Gly Arg Phe Gln Ala Gln Arg Leu Arg Arg Pro Pro Glu Asp 165 170 Arg Pro Asp Pro Leu Gly Thr Trp Leu Thr Glu Val Ala Gly Arg Phe Gly Val Glu Phe Gly Glu Asp Leu Ala Val Gly Gln Trp Ser Val Asp 200 Gln Leu Pro Pro Ser Phe Arg Leu Asp Thr Gly Met Glu Thr Val Val Ala Arg Thr Leu Pro Tyr Asn Gly Ala Ser Val Val Pro Asp Trp Leu Lys Lys Gly Ser Ala Thr Arg Arg Ile Cys Ile Thr Gly Gly Phe Ser Gly Leu Gly Leu Ala Ala Asp Ala Asp Gln Phe Ala Arg Thr Leu Ala Gln Leu Ala Arg Phe Asp Gly Glu Ile Val Val Thr Gly Ser Gly Pro Asp Thr Ser Ala Val Pro Asp Asn Ile Arg Leu Val Asp Phe Val Pro

315

320

Met Gly Val Leu Gln Asn Cys Ala Ala Ile Ile His His Gly Gly

Ala Gly Thr Trp Ala Thr Ala Leu His His Gly Ile Pro Gln Ile Ser 325 330 335

Val Ala His Glu Trp Asp Cys Met Leu Arg Gly Gln Gln Thr Ala Glu 340 345 350

Leu Gly Ala Gly Ile Tyr Leu Arg Pro Asp Glu Val Asp Ala Asp Ser 355 360 365

Leu Ala Ser Ala Leu Thr Gln Val Val Glu Asp Pro Thr Tyr Thr Glu 370 380

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Thr Gly Thr His Leu Arg Arg Phe Ala Asp Leu Phe Asp Asp Val Thr 50 55 60

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Arg Asp Gly Ala Glu Leu Asp Gln Ala Leu Ala Ser Phe Ala Arg His
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Leu Thr Ile Ser Arg Val Ser His Ser Val Arg Ala Gly Gly Ala Thr
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His His Val Glu His Tyr Glu Ile Thr Leu Phe Glu Arg Gln Gln Tyr 195 200 205

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Asp Arg Trp Ser Phe Gln Pro Glu Asp Gly Arg Leu Ala His Glu Ser 85 90 95

Gly Arg Phe Phe Ser Ile Glu Gly Leu His Val Arg Thr Asn Phe Gly 100 105 110

Trp Arg Arg Asp Trp Ile Gln Pro Ile Ile Val Gln Pro Glu Ile Gly
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Pro Thr Leu Gln Ala Thr Arg Ser Asn Tyr Thr Gly Val His Arg Gly
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Ser Lys Val Arg Phe Ile Glu Tyr Phe Asn Gly Thr Arg Pro Ser Arg 180 185 190

Ile Leu Val Asp Val Leu Gln Ser Glu Gln Gly Ala Trp Phe Leu Arg

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Thr Tyr Leu Leu Gly His Ser His Tyr Val Asn Ile Gln Leu Arg Ser 485

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Thr Ser Leu Tyr Gly Leu Asn Asn Asp Glu Tyr Glu Thr Thr Thr Gly

Gln Thr Leu Asn Arg Gln Pro Gly Leu Trp Glu Leu Gly Ala Ile Arg

Lys Arg Gly Ala Phe Glu Arg Val Lys Lys Asn Leu Gln Gly Phe Leu

260 265 270

Arg Met Arg Ala Glu Arg Asp Ala Pro Ile Arg Leu Gly Phe Asn His 275 280 285

Ile Ile Leu Pro Gly Arg Ala Asp Arg Leu Thr Asp Leu Val Asp Phe 290 295 300

Ile Ala Glu Leu Asn Glu Ser Ser Pro Gln Arg Pro Leu Asp Phe Val 305 310 315 320

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Glu Ser Leu Arg Arg Gly Val Asp Ala Glu Leu Leu Arg Ile Arg Pro 370 375 380

Glu Thr Met Arg Pro Thr Ala His Pro Gln Val Ala Val Gln Ile Asp 385 390 395 400

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Glu Gly Ala Thr Arg Tyr Ile Ala Gly Arg Val Thr Pro Ser Thr Ser 420 425 430

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Pro Arg Pro Gly Asp Glu Tyr Phe Leu Asp Gly Phe Asp Gln Ser Val 450 455 460

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		ttc Phe 100										336
		ttc Phe										384
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		acc Thr										672
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		_			_	_	gac Asp	_	_	_			_	_	960
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Asn Gly Gly Pro Leu Val Arg Glu Phe Glu Gly Arg Val Ala Asp Leu 50 55 60

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ctc	999	ctc	ctt	ccc	999	cag	tgg	ccc	gtg	ccc	tgc	acc	ggc	cgg	gtg	996

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579914\_1

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gag Glu																2636
ccc Pro	_			_	_	_	_	_	_	_	_					2684
cat (											gacg			gta Val 830		2733
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330

Pro Pro Ala Pro Ser Val Phe Gly Ala Ala Phe Glu Asn Ala Leu 345

340

Ala Glu Pro Leu Val Arg Ala Val Thr Gly Ala Ala Leu Gln Ala Leu Ala Glu Gly Pro Pro Arg Leu Thr Ala Ala Gly Pro Val Val Arg Arg 375 380 Arg Arg Ser Pro Val Val Gly Gly Leu His Arg Ala Pro Val Ala Ala 390 395 Ala <210> 17 <211> 426 <212> PRT <213> Streptomyces antibioticus <400> 17 Met Met Met Thr Thr Phe Ala Ala Asn Thr His Phe Gln Pro Leu Val Pro Leu Ala Trp Ala Leu Arg Thr Ala Gly His Glu Val Arg Val Val Ser Gln Pro Ser Leu Ser Asp Val Val Thr Gln Ala Gly Leu Thr Ser 40 Val Pro Val Gly Thr Glu Ala Pro Val Glu Gln Phe Ala Ala Thr Trp Gly Asp Asp Ala Tyr Ile Gly Val Asn Ser Ile Asp Phe Thr Gly Asn Asp Pro Gly Leu Trp Thr Trp Pro Tyr Leu Leu Gly Met Glu Thr Met Leu Val Pro Ala Phe Tyr Glu Leu Leu Asn Asn Glu Ser Phe Val Asp 105 Gly Val Val Glu Phe Ala Arg Asp Trp Arg Pro Asp Leu Val Ile Trp 115 Glu Pro Leu Thr Phe Ala Gly Ala Val Ala Ala Arq Val Thr Gly Ala Ala His Ala Arg Leu Pro Trp Gly Gln Glu Ile Thr Leu Arg Gly Arg Gln Ala Phe Leu Ala Glu Arg Ala Leu Gln Pro Phe Glu His Arg Glu

190

205

170

Asp Pro Thr Ala Glu Trp Leu Gly Arg Met Leu Asp Arg Tyr Gly Cys

Ser Phe Asp Glu Glu Met Val Thr Gly Gln Trp Thr Ile Asp Thr Leu

200

185

180

195

Pro Arg Ser Met Arg Leu Glu Leu Ser Glu Glu Leu Arg Thr Leu Asp 210 215 220

Met Arg Tyr Val Pro Tyr Asn Gly Pro Ala Val Val Pro Pro Trp Val 225 230 · 235 240

Trp Glu Pro Cys Glu Arg Pro Arg Val Cys Leu Thr Ile Gly Thr Ser 245 250 255

Gln Arg Asp Ser Gly Arg Asp His Val Pro Leu Asp His Leu Leu Asp 260 265 270

Ser Leu Ala Asp Val Asp Ala Glu Ile Val Ala Thr Leu Asp Thr Thr 275 280 285

Gln Glu Arg Leu Arg Gly Ala Ala Pro Gly Asn Val Arg Leu Val 290 295 300

Asp Phe Val Pro Leu His Ala Leu Met Pro Thr Cys Ser Ala Ile Val 305 310 315 320

His His Gly Gly Pro Gly Thr Trp Ser Thr Ala Ala Leu His Gly Val 325 330 335

Pro Gln Ile Ile Leu Asp Thr Ser Trp Asp Thr Pro Val Arg Ala Gln 340 345 350

Arg Met Gln Gln Leu Gly Ala Gly Leu Ser Met Pro Val Gly Glu Leu 355 . 360 . 365

Gly Val Glu Ala Leu Arg Asp Arg Val Leu Arg Leu Leu Gly Glu Pro 370 375 380

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His Ala Thr Gly Ala Met Ala Gly Arg Arg 420 425

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<212> PRT

<213> Streptomyces antibioticus

<400> 18

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20 25 30

Val Ala Ser Gln Pro Ala Leu Ser Asp Thr Ile Thr Gln Ala Gly Leu

Thr	Ala 50	Val	Pro	Val	Gly	Arg 55	Asp	Thr	Ala	Phe	Leu 60	Glu	Leu	Met	Gly
Glu 65	Ile	Gly	Ala	Asp	Val 70	Gln	Lys	Tyr	Ser	Thr 75	Gly	Ile	Asp	Leu	Gly 80
Val	Arg	Ala	Glu	Leu 85	Thr	Ser	Trp	Glu	Tyr 90	Leu	Leu	Gly	Met	His 95	Thr
Thr	Leu	Val	Pro 100	Thr	Phe	Tyr	Ser	Leu 105	Val	Asn	Asp	Glu	Pro 110	Phe	Val
Asp	Gly	Leu 115	Val	Ala	Leu	Thr	Arg 120	Ala	Trp	Arg	Pro	Asp 125	Leu	Ile	Leu
Trp	Glu 130	His	Phe	Ser	Phe	Ala 135	Gly	Ala	Leu	Ala	Ala 140	Arg	Ala	Thr	Gly
Thr 145	Pro	His	Ala	Arg	Val 150	Leu	Trp	Gly	Ser	Asp 155	Leu	Ile	Val	Arg	Phe 160
Arg	Arg	Asp	Phe	Leu 165	Ala	Glu	Arg	Ala	Asn 170	Arg	Pro	Ala	Glu	His 175	Arg
Glu	Asp	Pro	Met 180	Ala	Glu	Trp	Leu	Gly 185	Trp	Ala	Ala	Glu	Arg 190	Leu	Gly
Ser	Thr	Phe 195	Asp	Glu	Glu	Leu	Val 200	Thr	Gly	Gln	Trp	Thr 205	Ile	Asp	Pro
Leu	Pro 210	Arg	Ser	Met	Arg	Leu 215	Pro	Thr	Gly	Thr	Thr 220	Thr	Val	Pro	Met
Arg 225	Tyr	Val	Pro	Tyr	Asn 230	Gly	Arg	Ala	Val	Val 235	Pro	Ala	Trp	Val	Arg 240
Gln	Arg	Ala	Arg	Arg 245	Pro	Arg	Ile	Cys	Leu 250	Thr	Leu	Gly	Val	Ser 255	Ala
Arg	Gln	Thr	Leu 260	Gly	Asp	Gly	Val	Ser 265	Leu	Ala	Glu	Val	Leu 270	Ala	Ala
Leu	Gly	Asp 275	Val	Asp	Ala	Glu	Ile 280	Val	Ala	Thr	Leu	Asp 285	Ala	Ser	Gln
Arg	Lys 290	Leu	Leu	Gly	Pro	Val 295.		Asp	Asn	Val	Arg 300	Leu	Val	Asp	Phe
Val 305	Pro	Leu	His	Ala	Leu 310	Met	Pro	Thr	Cys	Ser 315	Ala	Ile	Val	His	His 320
Gly	Gly	Ala	Gly	Thr 325	Trp	Leu	Thr	Ala	Ala 330	Val	His	Glỳ	Val	Pro 335	Gln
Ile	Val	Leu	Gly	Asp	Leu	Trp	Asp	Asn	Leu	Leu	Arg	Ala	Arg	Gln	Thr

340 345 350

Gln Ala Ala Gly Ala Gly Leu Phe Ile His Pro Ser Glu Val Thr Ala 355 360 365

Ala Gly Leu Gly Glu Gly Val Arg Arg Val Leu Thr Asp Pro Ser Ile 370 375 380

Arg Ala Ala Gln Arg Val Arg Asp Glu Met Asn Ala Glu Pro Thr 385 390 395 400

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Arg Gly Arg Gly Gly Asn His Ala Gly
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<213> Streptomyces antibioticus

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Ala Glu Ile Thr Ala Ala Leu Val Asp Glu Leu Leu Phe Arg Cys Glu 35 40 45

Ile Pro Gln Val Gly Gly Glu Ala Phe Ile Gly Leu Asp Val Leu His 50 55 60

Gly Ala Asp Arg Ile Ser His Val Leu Gln Val Thr Asp Gly Lys Pro 65 70 75 80

Val Thr Ser Ala Glu Pro Ala Gly Gln Glu Leu Gly Gly Arg Thr Trp 85 90 95

Ser Ser Arg Ser Ala Thr Leu Leu Arg Glu Leu Phe Gly Pro Pro Ser

Gly Arg Thr Ala Gly Gly Phe Gly Val Ser Phe Leu Pro Asp Leu Arg 115 120 125

Gly Pro Arg Thr Met Glu Gly Ala Ala Leu Ala Ala Arg Ala Thr Asn 130 135 140

Val Val Leu His Ala Thr Thr Asn Glu Thr Pro Pro Leu Asp Arg Leu 145 150 155 160

Ala Leu Arg Tyr Glu Ser Asp Lys Trp Gly Gly Val His Trp Phe Thr 165 170 175

Gly His Tyr Asp Arg His Leu Arg Ala Val Arg Asp Gln Ala Val Arg Ile Leu Glu Ile Gly Ile Gly Gly Tyr Asp Asp Leu Leu Pro Ser Gly 200 Ala Ser Leu Lys Met Trp Lys Arg Tyr Phe Pro Arg Gly Leu Val Phe 215 Gly Val Asp Ile Phe Asp Ser Arg Arg Ala Thr Ser Arg Val Ser Arg 230 235 Arg Ser Ala Ala Arg Gln Asp Asp Pro Glu Phe Met Arg Arg Val Ala 250 Glu Glu His Gly Pro Phe Asp Val Ile Ile Asp Asp Gly Ser His Ile Asn Ala His Met Arg Thr Ser Phe Ser Val Met Phe Pro His Leu Arg Asn Gly Gly Phe Tyr Val Ile Glu Asp Thr Phe Thr Ser Tyr Trp Pro 295 Gly Tyr Gly Gly Pro Ser Gly Ala Arg Cys Pro Ser Gly Thr Thr Ala Leu Glu Met Val Lys Gly Leu Ile Asp Ser Val His Tyr Glu Glu Arg 330 Pro Asp Gly Ala Ala Thr Ala Asp Tyr Ile Ala Arg Asn Leu Val Gly Leu His Ala Tyr Gln Thr Thr Ser Ser Ser Arg Arg Ala Ile Asn 360 Lys Glu Gly Gly Ile Pro His Thr Val Pro Arg Glu Pro Phe Trp Asn 375 Asp Asn 385 <210> 20 <211> 738 <212> DNA <213> Streptomyces antibioticus <220> <221> CDS <222> (1)..(738) <223> /gene= "oleM" /note= "SEQ ID No. 15 from 3992 to 4729" <220> <221> mat\_peptide <222> (1)

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			_	gac Asp		_	_		_				_			192
	_	_		ttc Phe	_	_	_	_			_	_			_	240
_	_	_		gcc Ala 85	_		_	_	_			_	-			288
				tcc Ser												336
				tcc Ser												384
				tgc Cys												432
_		-	_	tgg Trp			_					_				480
				gta Val 165												528
				gac Asp												576
				gag Glu												624
	_	_		ccg Pro					_	_				_		672

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Glu Ala Lys Asp 35	Val Ala Asp	Leu Val Ar 40		l Pro Asp 5	Ala
Ser Ser Leu Leu 50	Asp Val Ala		nr Gly Ala Hi 60	s Leu Arg	His
Phe Ala Thr Leu 65	Phe Asp Asp 70	Ala Arg Gl	ly Leu Glu Le 75	u Ser Ala	Ser 80
Met Leu Asp Ile	Ala Arg Sen 85	•	ro Gly Val Pr 90	o Leu His 95	Gln
Gly Asp Met Arg		Leu Gly Pr 105	ro Arg Val Se	er Ala Val 110	Thr
Cys Met Phe Ser 115	Ser Val Gly	His Leu Al	la Thr Thr Al		Asp
Ala Thr Leu Arg	Cys Phe Ala	_	nr Arg Pro Gl 140	y Gly Val	Ala
Val Ile Glu Pro 145	Trp Trp Phe	e Pro Glu Th	nr Phe Thr As 155	sp Gly Tyr	Val 160
Ala Gly Asp Ile	Val Arg Val		rg Thr Ile Se 70	er Arg Val 175	Ser
His Ser Val Arg		/ Ala Thr Ai 185	rg Met Glu Il	e His Tyr 190	Val
Ile Ala Asp Ala 195	Glu His Gly	Pro Arg Hi	is Leu Val Gl 20		Arg
Ile Thr Leu Phe 210	Pro Arg His		nr Ala Ala Ty 220	r Glu Lys	Ala

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225
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gacgaggccg aagaggtgg
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ccgtcgagct ctgaggtaa
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tgcacgcgct gctgccgacc
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- 55 **-**

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- G - C

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B' cont